

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

#### TABLE OF CONTENTS

1. REAL PARTY IN INTEREST	2
2. RELATED APPEALS AND INTERFERENCES	2
3. STATUS OF CLAIMS	2
4. STATUS OF AMENDMENTS	2
5. SUMMARY OF THE CLAIMED SUBJECT MATTER	2
6. GROUNDS OF REJECTION TO BE REVIEWED ON APPEAL	6
7. ARGUMENT	6
8. CLAIM APPENDIX	Attached
9. EVIDENCE APPENDIX	Attached
10. RELATED PROCEEDINGS APPENDIX	None

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE  
BEFORE THE BOARD OF PATENT APPEALS AND INTERFERENCES

Application No.: 10/052,798  
Applicant: Adams, et al.  
Filed: November 2, 2001  
Group Art Unit: 1646  
Examiner: Eileen O'Hara  
Docket No.: 22338-00904/P1101R2D1

**APPEAL BRIEF**

Mail Stop Appeal Brief-Patents  
Commissioner for Patents  
P.O. Box 1450  
Washington, D.C. 20231

Dear Sir:

A Notice of Appeal was filed on September 16, 2005. A four-month Extension of Time was previously filed with a response on March 15, 2006. Accordingly, Applicant submits that this Appeal Brief filed under 37 CFR § 41.37, which is being filed with an additional one-month Extension of Time, is timely filed. Appellant requests the Commissioner to charge Deposit Account No. 18-1260 for the \$500.00 Appeal Brief fee due under 37 CFR 41.20(b)(2).

04/18/2006 TL0111 00000076 181260 10052798  
02 FC:1402 500.00 DA

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

#### **1. REAL PARTY IN INTEREST**

The real party in interest in this appeal is Genentech, Inc.

#### **2. RELATED APPEALS AND INTERFERENCES**

The appeals or interferences known to the Appellant, the Appellant's legal representative, or assignee which may be related to, directly affect or be directly affected by or to have a bearing on the Board's decision in the pending appeal may include the following:

Patent Interference Nos: 105,361; 105,240; 105,380; and 105,381.

#### **3. STATUS OF CLAIMS**

Claims 1-64 were originally filed. Claims 65-97 were added, claims 59-62 were amended and original claims 1-58 and 63-64 were cancelled at the time of filing in a preliminary amendment. Claims 98-146 were added and entered into the application during prosecution. *See, e.g.,* Advisory Action dated November 23, 2005. The claims involved in this appeal, claims 59-62 and 64-146, are presented in the claim appendix attached hereto.

#### **4. STATUS OF AMENDMENTS**

There are no outstanding amendments.

#### **5. SUMMARY OF CLAIMED SUBJECT MATTER**

- In one aspect, the present invention relates to methods of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cancer cell *in vivo* or *ex vivo*. *See, e.g.,* SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

- In another aspect, the present invention relates to methods of treating mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cancer cell *in vivo* or *ex vivo*. *See, e.g.*, SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 54 to 182 SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. *See, e.g.*, SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 1 to 182 of SEQ ID:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. *See, e.g.*, SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of treating mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

polypeptide consisting of amino acids 54 to 182 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.

- In another aspect, the present invention relates to methods of treating mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 1 to 182 of SEQ ID NO: 1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of inducing apoptosis in mammalian cells expressing Apo-2 receptor comprising exposing mammalian cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody, wherein said antibody stimulates apoptosis in said mammalian cells upon its binding to said Apo-2 receptor, and wherein said antibody binds to an Apo-2 receptor polypeptide having at least about 80% sequence identity to SEQ ID NO:1. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of treating cancer, comprising exposing mammalian cancer cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody, wherein said antibody stimulates apoptosis in said cells upon its binding to said Apo-2 receptor, and wherein said antibody binds to an Apo-2 receptor polypeptide having at least about 80% sequence identity to SEQ ID NO:1. See, e.g., SEQ

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.

- In another aspect, the present invention relates to methods of inducing apoptosis in mammalian cells expressing Apo-2 receptor comprising exposing mammalian cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of inducing apoptosis in mammalian cells expressing Apo-2 receptor comprising exposing mammalian cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide which consists of amino acid residues 54 to 182 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.
- In another aspect, the present invention relates to methods of treating cancer, comprising exposing mammalian cancer cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cancer cell *in vivo* or *ex vivo*. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

- In another aspect, the present invention relates to methods of treating cancer, comprising exposing mammalian cancer cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide which consists of amino acid residues 54 to 182 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*. See, e.g., SEQ ID NO:1; Figures 1 to 16; pages 12, line 37 to page 14, line 36; page 16, line 21 to page 19, line 14; page 34, lines 28 to page 35, line 26; page 47, lines 24 to page 61; pages 62 to 88.

## 6. GROUNDS OF REJECTION TO BE REVIEWED ON APPEAL

Whether the provisional rejection of pending claims 59-62, 65-75, 79-89 and 93-97 (and the objection to claims 76-78 and 90-92 as depending thereon) for obviousness-type double patenting over claims 1 to 5 and 10 to 47 of U.S.S.N. 09/396,710 ("the '710 application") is proper.

## 7. ARGUMENT

The only rejection pending in the application on appeal is a provisional double patenting rejection based on claims that were pending, prior to April 14, 2006, in the '710 application. No other rejection has been maintained by the Examiner.

On April 14, 2006, the Appellant expressly abandoned the '710 application. A copy of the express abandonment and its proof of filing is provided in the Evidence Appendix.

As noted on the letter expressly abandoning the '710 application, the express abandonment of the '710 application is not an abandonment of subject matter contained in that application. Moreover, Appellant notes that a continuation application claiming the benefit of the applications to which the '710 application claims benefit under 35 U.S.C. §120 was filed on April 13, 2006. The April 13, 2006 continuation contained a preliminary amendment cancelling all original claims and presenting new claims. None of the new claims added by the April 13,

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

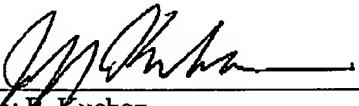
2006 preliminary amendment correspond to the claims previously pending in the '710 application. In addition, the claims added by the April 13, 2006, preliminary amendment correspond to claims previously held by the Examiner to be drawn to a patentably distinct invention relative to the claims that are the subject of the present appeal.

The abandonment of the '710 application and the absence of any other pending application containing claims that correspond to the '710 application removes the basis of the provisional double-patenting rejection of the claims pending in the application on appeal. As such, the only ground for rejection of the appealed claims is moot. It is respectfully submitted that the provisional rejection should be withdrawn and the pending claims passed to issue.

In the unlikely event that the transmittal letter is separated from this document and the Patent Office determines that an extension and/or other relief is required, Appellant petitions for any required relief including extensions of time and authorizes the Assistant Commissioner to charge the cost of such petitions and/or other fees due in connection with the filing of this document to Deposit Account No. 18-1260 referencing docket no. 2233800904. However, the Assistant Commissioner is not authorized to charge the cost of the issue fee to the Deposit Account.

Respectfully submitted,

Dated: April 17, 2006

By:   
Jeffrey P. Kushan  
Registration No. 43,401  
Attorney for Appellant

Sidley Austin LLP  
1501 K Street, NW  
Washington, DC 20005  
Phone: 202-736-8157  
Fax: 202-736-8711

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

## 8. CLAIM APPENDIX

59. (previously presented) A method of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cancer cell *in vivo* or *ex vivo*.

60. (previously presented) The method of claim 59 wherein said antibody comprises a single-chain antibody.

61. (previously presented) A method of treating mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cancer cell *in vivo* or *ex vivo*.

62. (previously presented) The method of claim 61 wherein said antibody comprises a single-chain antibody.

65. (previously presented) A method of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 54 to 182 SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.

66. (previously presented) A method of inducing apoptosis in mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 1 to 182 of SEQ ID:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

67. (previously presented) A method of treating mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 54 to 182 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.

68. (previously presented) A method of treating mammalian cancer cells comprising exposing mammalian cancer cells to an effective amount of monoclonal antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide consisting of amino acids 1 to 182 of SEQ ID NO: 1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.

69. (previously presented) The method of claim 59, 65, or 66, wherein said antibody is a chimeric antibody.

70. (previously presented) The method of claim 59, 65, or 66, wherein said antibody is a humanized antibody.

71. (previously presented) The method of claim 59, 65, or 66, wherein said antibody is a human antibody.

72. (previously presented) The method of claim 59, 65, or 66, wherein said antibody comprises an Fab fragment.

73. (previously presented) The method of claim 59, 65, or 66, wherein said antibody comprises a scFv fragment.

74. (previously presented) The method of claim 59, 65, or 66, wherein said antibody comprises a F(ab')2 fragment.

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

75. (previously presented) The method of claim 59, 65, or 66, wherein said antibody binds to the same epitope as the epitope to which the monoclonal antibody produced by the hybridoma cell line deposited as ATCC accession number HB-12456 binds.

76. (previously presented) The method of claim 59, 65, or 66, wherein said antibody comprises one or more complementary determining region (CDR) amino acid sequences of the 16E2 antibody shown in Figure 16 (SEQ ID NO:9).

77. (previously presented) The method of claim 59, 65, or 66, wherein said antibody comprises one or more complementary determining region (CDR) amino acid sequences of the 20E6 antibody shown in Figure 16 (SEQ ID NO:10).

78. (previously presented) The method of claim 59, 65, or 66, wherein said antibody comprises one or more complementary determining region (CDR) amino acid sequences of 24C4 antibody shown in Figure 16 (SEQ ID NO:11).

79. (previously presented) The method of claim 59, 65, or 66, wherein said antibody is fused to an epitope tag sequence.

80. (previously presented) The method of claim 59, 65, or 66, wherein the cancer cells are colon or colorectal cancer cells.

81. (previously presented) The method of claim 59, 65, or 66, wherein the cancer cells are lung cancer cells.

82. (previously presented) The method of claim 59, 65, or 66, wherein the cancer cells are breast cancer cells.

83. (previously presented) The method of claim 61, 67, or 68, wherein said antibody is a chimeric antibody.

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

84. (previously presented) The method of claim 61, 67, or 68, wherein said antibody is a humanized antibody.
85. (previously presented) The method of claim 61, 67, or 68, wherein said antibody is a human antibody.
86. (previously presented) The method of claim 61, 67, or 68, wherein said antibody comprises an Fab fragment.
87. (previously presented) The method of claim 61, 67, or 68, wherein said antibody comprises a scFv fragment.
88. (previously presented) The method of claim 61, 67, or 68, wherein said antibody comprises a F(ab') 2 fragment.
89. (previously presented) The method of claim 61, 67, or 68, wherein said antibody binds to the same epitope as the epitope to which the monoclonal antibody produced by the hybridoma cell line deposited as ATCC accession number HB-12456 binds.
90. (previously presented) The method of claim 61, 67, or 68, wherein said antibody comprises one or more complementary determining region (CDR) amino acid sequences of the 16E2 antibody shown in Figure 16 (SEQ ID NO:9).
91. (previously presented) The method of claim 61, 67, or 68, wherein said antibody comprises one or more complementary determining region (CDR) amino acid sequences of the 20E6 antibody shown in Figure 16 (SEQ ID NO:10).
92. (previously presented) The method of claim 61, 67, or 68, wherein said antibody comprises one or more complementary determining region (CDR) amino acid sequences of the 24C4 antibody shown in Figure 16 (SEQ ID NO:11).

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

93. (previously presented) The method of claim 61, 67, or 68, wherein said antibody is fused to an epitope tag sequence.

94. (previously presented) The method of claim 61, 67, or 68, wherein said mammalian cancer cells are exposed to chemotherapy or radiation therapy.

95. (previously presented) The method of claim 61, 67, or 68, wherein the cancer cells are colon or colorectal cancer cells.

96. (previously presented) The method of claim 61, 67, or 68, wherein the cancer cells are lung cancer cells.

97. (previously presented) The method of claim 61, 67, or 68, wherein the cancer cells are breast cancer cells.

98. (New) A method of inducing apoptosis in mammalian cells expressing Apo-2 receptor comprising exposing mammalian cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody, wherein said antibody stimulates apoptosis in said mammalian cells upon its binding to said Apo-2 receptor, and wherein said antibody binds to an Apo-2 receptor polypeptide having at least about 80% sequence identity to SEQ ID NO:1.

99. (New) The method of claim 98 wherein said Apo-2 agonist antibody is a monoclonal antibody.

100. (New) The method of claim 98 wherein said agonist antibody is a chimeric antibody.

101. (New) The method of claim 98 wherein said agonist antibody is a humanized antibody.

102. (New) The method of claim 98 wherein said agonist antibody is a human antibody.

103. (New) A method of treating cancer, comprising exposing mammalian cancer cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody, wherein said antibody stimulates apoptosis in said cells upon its binding to said Apo-2 receptor, and wherein

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

said antibody binds to an Apo-2 receptor polypeptide having at least about 80% sequence identity to SEQ ID NO:1.

104. (New) The method of claim 103, wherein said cancer cells are lung cancer cells.
105. (Original) The method of claim 103, wherein said cancer cells are colon cancer cells.
106. (Original) The method of claim 103, wherein said cancer cells are glioma cells.
107. (New) A method of inducing apoptosis in mammalian cells expressing Apo-2 receptor comprising exposing mammalian cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.
108. (New) The method of claim 107 wherein said Apo-2 agonist antibody is a monoclonal antibody.
109. (New) The method of claim 107 wherein said agonist antibody is a chimeric antibody.
110. (New) The method of claim 107 wherein said agonist antibody is a humanized antibody.
111. (New) The method of claim 107 wherein said agonist antibody is a human antibody.
112. (New) The method of claim 107 wherein said mammalian cells expressing Apo-2 receptor are cancer cells.
113. (New) The method of claim 112 wherein said cancer cells are lung cancer cells.
114. (New) The method of claim 112 wherein said cancer cells are colon cancer cells.
115. (New) The method of claim 112 wherein said cancer cells are glioma cells.

839396v1<DCI>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

116. (New) A method of inducing apoptosis in mammalian cells expressing Apo-2 receptor comprising exposing mammalian cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide which consists of amino acid residues 54 to 182 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.
117. (New) The method of claim 116 wherein said Apo-2 agonist antibody is a monoclonal antibody.
118. (New) The method of claim 116 wherein said agonist antibody is a chimeric antibody.
119. (New) The method of claim 116 wherein said agonist antibody is a humanized antibody.
120. (New) The method of claim 116 wherein said agonist antibody is a human antibody.
121. (New) The method of claim 116 wherein said mammalian cells expressing Apo-2 receptor are cancer cells.
122. (New) The method of claim 121 wherein said cancer cells are lung cancer cells.
123. (New) The method of claim 121 wherein said cancer cells are colon cancer cells.
124. (New) The method of claim 121 wherein said cancer cells are glioma cells.
125. (New) A method of treating cancer, comprising exposing mammalian cancer cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to Apo-2 polypeptide consisting of the contiguous amino acid residues 1 to 411 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cancer cell *in vivo* or *ex vivo*.
126. (New) The method of claim 125 wherein said Apo-2 agonist antibody is a monoclonal antibody.

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

127. (New) The method of claim 125 wherein said agonist antibody is a chimeric antibody.
128. (New) The method of claim 125 wherein said agonist antibody is a humanized antibody.
129. (New) The method of claim 125 wherein said agonist antibody is a human antibody.
130. (New) The method of claim 125 wherein said mammalian cancer cells are lung cancer cells.
131. (New) The method of claim 125 wherein said mammalian cancer cells are colon cancer cells.
132. (New) The method of claim 125 wherein said mammalian cancer cells are glioma cells.
133. (New) A method of treating cancer, comprising exposing mammalian cancer cells expressing Apo-2 receptor to an effective amount of an Apo-2 agonist antibody which (a) binds to a soluble extracellular domain sequence of an Apo-2 polypeptide which consists of amino acid residues 54 to 182 of SEQ ID NO:1 and (b) stimulates apoptosis in at least one type of mammalian cell *in vivo* or *ex vivo*.
134. (New) The method of claim 133 wherein said Apo-2 agonist antibody is a monoclonal antibody.
135. (New) The method of claim 133 wherein said agonist antibody is a chimeric antibody.
136. (New) The method of claim 133 wherein said agonist antibody is a humanized antibody.
137. (New) The method of claim 133 wherein said agonist antibody is a human antibody.
138. (New) The method of claim 133 wherein said mammalian cancer cells are lung cancer cells.
139. (New) The method of claim 133 wherein said mammalian cancer cells are colon cancer cells.

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

140. (New) The method of claim 133 wherein said mammalian cancer cells are glioma cells.
141. (New) The method of claim 98 wherein said Apo-2 receptor polypeptide has at least about 85% sequence identity to SEQ ID NO:1.
142. (New) The method of claim 98 wherein said Apo-2 receptor polypeptide has at least about 90% sequence identity to SEQ ID NO:1.
143. (New) The method of claim 98 wherein said Apo-2 receptor polypeptide has at least about 95% sequence identity to SEQ ID NO:1.
144. (New) The method of claim 103 wherein said Apo-2 receptor polypeptide has at least about 85% sequence identity to SEQ ID NO:1.
145. (New) The method of claim 103 wherein said Apo-2 receptor polypeptide has at least about 90% sequence identity to SEQ ID NO:1.
146. (New) The method of claim 103 wherein said Apo-2 receptor polypeptide has at least about 95% sequence identity to SEQ ID NO:1.

839396v1<DC1>

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

**9. EVIDENCE APPENDIX**

- Copy of Express Abandonment filed in U.S.S.N. 09/396,710 with fax filing receipt

839396v1<DCI>

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Application of  Avi J. Ashkenazi  Confirmation No. 7837  Serial No.: 09/396,710  Filed: September 15, 1999  For: APO-2 RECEPTOR ANTIBODIES	Patent Docket P1101P2/22338-00901  Group Art Unit: 1646  Examiner: Claire M. Kaufman
--	--

## EXPRESS ABANDONMENT UNDER 37 CFR 1.138

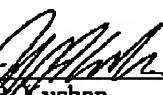
Commissioner of Patents  
United States Patent and Trademark Office  
P.O. Box 1450  
Alexandria, VA 22313-1450

Sir:

Applicant herewith expressly abandons the above-captioned application, without disclaimer of the right of the Applicant to present and secure in the future claims to subject matter that corresponds or is related to the claims of this application. The express abandonment of this application is not being made in response to any pending rejection of the subject matter of the claims. Applicant further observes that this express abandonment is made concurrently with the filing of a continuation application of U.S.S.N. 09/396,710.

Respectfully submitted,

Date: April 14, 2006

By:   
Jeffrey P. Kushan  
Reg. No. 43,401  
Attorney for Applicant

SIDLEY AUSTIN LLP  
1501 K Street, N.W.  
Washington, DC 20005  
202-736-8914 (direct)  
202-736-8000 (main)  
202-736-8711 (fax)

USPTO  
TO:Auto-reply fax to 202 736 8714 COMPANY:

4/14/2006 9:04 AM

PAGE 1/001 Fax Server

## Auto-Reply Facsimile Transmission



TO:

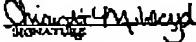
Fax Information  
Date Received:  
Total Pages:

Fax Sender at 202 736 8714

4/14/2006 9:01:59 AM [Eastern Daylight Time]  
2 (including cover page)

**ADVISORY:** This is an automatically generated return receipt confirmation of the facsimile transmission received by the Office. Please check to make sure that the number of pages listed as received in Total Pages above matches what was intended to be sent. Applicants are advised to retain this receipt in the unlikely event that proof of this facsimile transmission is necessary. Applicants are also advised to use the certificate of facsimile transmission procedures set forth in 37 CFR 1.8(a) and (b), 37 CFR 1.6(f). Trademark Applicants, also see the Trademark Manual of Examining Procedure (TMEP) section 306 et seq.

Received  
Cover  
Page  
=====>

04/14/06 09:01 FAX 202 736 8714 1A-LLP-DC		022
<b>SIDLEY AUSTIN LLP</b> 1250 K STREET, N.W. - WASHINGTON, D.C. 20004 • TEL. (202) 736-6000 • FAX (202) 736-6714		
<b>FAX TRANSMISSION</b>		
Total <b>2</b> / pages, including cover sheet To: Commissioner for Patents U.S. Patent and Trademark Office Jeffrey P. Kuskin Tel. (202) 736-6000 Date: April 14, 2006		
From: (571) 273-8300 Examiner: Clive M. KAUFMAN		
Re: Serial No.: 09/396,710 Group Art Unit: 1646 Filed: September 15, 1999 Applicant: Avi A. ASHKENAZI For: APO-2 RECEPTOR ANTIODIES		
<b>CERTIFICATE OF TRANSMISSION UNDER 37 C.F.R. § 1.8</b> I CERTIFY THAT THE FOLLOWING DOCUMENTS ARE BEING TRANSMITTED TO THE CPTO AT FAX NUMBER (571) 273-8300 ON THE DATE SHOWN. 1. Express Amendment Under 37 CFR, L131		
 Clifford M. Lloyd SIGNATURE		Clifford M. Lloyd PRINTED NAME
		April 14, 2006
<small>           THIS MESSAGE IS PROVIDED ONLY FOR THE USE OF THE ADDRESSEES OR ENTITY TO WHICH IT IS ADDRESSED AND MAY CONTAIN INFORMATION THAT IS PROPRIETARY, UNPUBLISHED OR OTHERWISE UNDUE UNDER APPLICABLE LAW. IF THE RECIPIENT OF THIS MESSAGE IS NOT THE INTENDED RECIPIENT OR HIS EMPLOYEE OR ASSISTANT, HE SHOULD NOTIFY THE SENDER SO THE INFORMATION IS NOT USED. THE RECIPIENT IS NOTIFIED THAT THIS INFORMATION, UNLESS OTHERWISE PROVIDED IN THE COMMUNICATION, IS ENTIRELY CONFIDENTIAL. IF YOU HAVE RECEIVED THIS INFORMATION IN ERROR, NOTIFY OR DISMISSE IT IMMEDIATELY BY TELEPHONE AND RETURN THE ORIGINAL MESSAGE TO US AT THE ADDRESS INDICATED VIA THE US MAIL OR SERVICE MARKS.         </small>		
<small>           PAGE 01 * PCDU AT 4/14/2006 9:01:59 AM [Eastern Daylight Time] * INQUIRIES@USPTO.GOV * CSID:202 736 8714 * DURATION (mm:ss):06:10         </small>		

04/14/06 FRI 09:03 [TX/RX NO 6043]

In Re Adams, et al.  
Application No. 10/052,798  
Appeal Brief

**10. RELATED PROCEEDINGS APPENDIX**

None

839396v1<DCI>

**This Page is Inserted by IFW Indexing and Scanning  
Operations and is not part of the Official Record**

## **BEST AVAILABLE IMAGES**

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images include but are not limited to the items checked:

- BLACK BORDERS**
- IMAGE CUT OFF AT TOP, BOTTOM OR SIDES**
- FADED TEXT OR DRAWING**
- BLURRED OR ILLEGIBLE TEXT OR DRAWING**
- SKEWED/SLANTED IMAGES**
- COLOR OR BLACK AND WHITE PHOTOGRAPHS**
- GRAY SCALE DOCUMENTS**
- LINES OR MARKS ON ORIGINAL DOCUMENT**
- REFERENCE(S) OR EXHIBIT(S) SUBMITTED ARE POOR QUALITY**
- OTHER:** \_\_\_\_\_

**IMAGES ARE BEST AVAILABLE COPY.**

**As rescanning these documents will not correct the image problems checked, please do not report these problems to the IFW Image Problem Mailbox.**